



From the INTERNATIONAL SE	ARCHING AUTHORILI	PUI
To: WOLF, GREENFIELD & SA Attn. Gates, Edward 600 Atlantic Avenue Boston, Massachusett UNITED STATES OF AME	R . File Folder ECB 57 Docket Entry Docket Cross	dt F Comme (DCT Dute AA 1)
	DOCKETED	Date of mailing (day/month/year) 28/03/2002
Applicant's or agent's file reference B0801/7228W0	APR 2 2002	FOR FURTHER ACTION See paragraphs 1 and 4 below
International application No. PCT/US 01/29636		International filing date (day/month/year) 21/09/2001
Applicant THE BRIGHAM AND WOME	N'S HOSPITAL, INC.	
Filing of amendments ar The applicant is entitled, if When? The time limit for International Sea Where? Directly to the For more detailed instruct The applicant is hereby no Article 17(2)(a) to that effe	Ind statement under Article 19 If he so wishes, to amend the clar filing such amendments is not arch Report; however, for more International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Fascimile No.: (41–22) 740.14 ctions, see the notes on the activitied that no International Seated is transmitted herewith.	aims of the International Application (see Rule 46): rmally 2 months from the date of transmittal of the details, see the notes on the accompanying sheet. 35 ccompanying sheet. arch Report will be established and that the declaration under
the protest together v	with the decision thereon has b	ditional fee(s) under Rule 40.2, the applicant is notified that: seen transmitted to the International Bureau together with the protest and the decision thereon to the designated Offices.
		applicant will be notified as soon as a decision is made.
Shortly after 18 months from the lifthe applicant wishes to avoid priority claim, must reach the lifth completion of the technical presentation. Within 19 months from the priority wishes to postpone the entry in the Within 20 months from the priority before all designated Offices with the priority in	d or postpone publication, a no International Bureau as provide eparations for international pub- rity date, a demand for internation into the national phase until 30 rity date, the applicant must per	I application will be published by the International Bureau. tice of withdrawal of the international application, or of the ed in Rules 90bis.1 and 90bis.3, respectively, before the lication. ional preliminary examination must be filed if the applicant months from the priority date (in some Offices even later).
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Name and mailing address of the International Searching Authority European Patent Office, P.B. 5818 Patentlaan 2 NL-2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016

Sandrine Parriche

Authorized officer

NOTES TO FORM PCT/ISA/220

These Notes are intended to give the basic instructions concerning the filing of amendments under article 19. The Notes are based on the requirements of the Patent Cooperation Treaty, the Regulations and the Administrative Instructions under that Treaty. In case of discrepancy between these Notes and those requirements, the latter are applicable. For more detailed information, see also the PCT Applicant's Guide, a publication of WIPO.

In these Notes, "Article", "Rule", and "Section" refer to the provisions of the PCT, the PCT Regulations and the PCT Administrative Instructions respectively.

INSTRUCTIONS CONCERNING AMENDMENTS UNDER ARTICLE 19

The applicant has, after having received the international search report, one opportunity to amend the claims of the international application. It should however be emphasized that, since all parts of the international application (claims, description and drawings) may be amended during the international preliminary examination procedure, there is usually no need to file amendments of the claims under Article 19 except where, e.g. the applicant wants the latter to be published for the purposes of provisional protection or has another reason for amending the claims before international publication. Furthermore, it should be emphasized that provisional protection is available in some States only.

What parts of the international application may be amended?

Under Article 19, only the claims may be amended.

During the international phase, the claims may also be amended (or further amended) under Article 34 before the International Preliminary Examining Authority. The description and drawings may only be amended under Article 34 before the International Examining Authority.

Upon entry into the national phase, all parts of the international application may be amended under Article 28 or, where applicable, Article 41.

When?

Within 2 months from the date of transmittal of the international search report or 16 months from the priority date, whichever time limit expires later. It should be noted, however, that the amendments will be considered as having been received on time if they are received by the International Bureau after the expiration of the applicable time limit but before the completion of the technical preparations for international publication (Rule 46.1).

Where not to file the amendments?

The amendments may only be filed with the International Bureau and not with the receiving Office or the International Searching Authority (Rule 46.2).

Where a demand for international preliminary examination has been its filed, see below.

How?

Either by cancelling one or more entire claims, by adding one or more new claims or by amending the text of one or more of the claims as filed.

A replacement sheet must be submitted for each sheet of the claims which, on account of an amendment or amendments, differs from the sheet originally filed.

All the claims appearing on a replacement sheet must be numbered in Arabic numerals. Where a claim is cancelled, no renumbering of the other claims is required. In all cases where claims are renumbered, they must be renumbered consecutively (Administrative Instructions, Section 205(b)).

The amendments must be made in the language in which the international application is to be published.

What documents must/may accompany the amendments?

Letter (Section 205(b)):

The amendments must be submitted with a letter.

The letter will not be published with the international application and the amended claims. It should not be confused with the "Statement under Article 19(1)" (see below, under "Statement under Article 19(1)").

The letter must be in English or French, at the choice of the applicant. However, if the language of the international application is English, the letter must be in English; if the language of the international application is French, the letter must be in French.

NOTES TO FORM PCT/ISA/220 (continued)

The letter must indicate the differences between the claims as filed and the claims as amended. It must, in particular, indicate, in connection with each claim appearing in the international application (it being understood that identical indications concerning several claims may be grouped), whether

- (i) the claim is unchanged;
- (ii) the claim is cancelled;
- (iii) the claim is new:
- (iv) the claim replaces one or more claims as filed;
- (v) the claim is the result of the division of a claim as filed.

The following examples illustrate the manner in which amendments must be explained in the accompanying letter:

- (Where originally there were 48 claims and after amendment of some claims there are 51):
 *Claims 1 to 29, 31, 32, 34, 35, 37 to 48 replaced by amended claims bearing the same numbers;
 claims 30, 33 and 36 unchanged; new claims 49 to 51 added."
- 2 [Where originally there were 15 claims and after amendment of all claims there are 11]: "Claims 1 to 15 replaced by amended claims 1 to 11."
- 3 [Where originally there were 14 claims and the amendments consist in cancelling some claims and in adding new claims! "Claims 1 to 6 and 14 unchanged; claims 7 to 13 cancelled; new claims 15, 16 and 17 added." or "Claims 7 to 13 cancelled; new claims 15, 16 and 17 added; all other claims unchanged."
- 4 [Where various tunds of amendments are made]:
 "Claims 1-10 unchanged; claims 11 to 13, 18 and 19 cancelled; claims 14, 15 and 16 replaced by amended claims 14, claim 17 subdivided into amended claims 15, 16 and 17; new claims 20 and 21 added."

"Statement under article 19(1)" (Rule 46.4)

The amendments may be accompanied by a statement explaining the amendments and indicating any impact that such amendments might have on the description and the drawings (which cannot be amended under Article 19(1))

The statement will be published with the international application and the amended claims.

It must be in the language in which the international appplication is to be published.

It must be brief, not exceeding 500 words if in English or if translated into English.

It should not be confused with and does not replace the letter indicating the differences between the claims as filed and as amended. It must be filed on a separate sheet and must be identified as such by a heading, preferably by using the words "Statement under Article 19(1)."

It may not contain any disparaging comments on the international search report or the relevance of citations contained in that report. Reference to citations, relevant to a given claim, contained in the international search report may be made only in connection with an amendment of that claim.

Consequence if a demand for international preliminary examination has already been filed

If, at the time of filing any amendments under Article 19, a demand for international preliminary examination has already been submitted, the applicant must preferably, at the same time of filing the amendments with the International Bureau, also file a copy of such amendments with the International Preliminary Examining Authority (see Rule 62.2(a), first sentence).

Consequence with regard to translation of the international application for entry into the national phase

The applicant's attention is drawn to the fact that, where upon entry into the national phase, a translation of the claims as amended under Article 19 may have to be furnished to the designated/elected Offices, instead of, or in addition to, the translation of the claims as filed.

For further details on the requirements of each designated/elected Office, see Volume II of the PCT Applicant's Guide.

PATENT COOPERATION TREATY

PCT

INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference	FOR FURTHER see Notification of Transmittal of International Search Report (Form PCT/ISA/220) as well as, where applicable, item 5 below.		
B0801/7228W0	International filing date (day/month/year)	(Earliest) Priority Date (day/month/year)	
International application No.			
PCT/US 01/29636	21/09/2001	21/09/2000	
THE BRIGHAM AND WOMEN'S H	OSPITAL, INC.		
according to Article 18. A copy is being tra This international Search Report consists	•		
1. Basis of the report			
	international search was carried out on the bas ess otherwise indicated under this item.	sis of the international application in the	
the international search w. Authority (Rule 23.1(b)).	as carried out on the basis of a translation of th	ne international application furnished to this	
was carried out on the basis of the contained in the internatio filed together with the inter furnished subsequently to furnished subsequently to the statement that the sub international application as	e sequence listing: nal application in written form. rnational application in computer readable form this Authority in written form. this Authority in computer readble form. rsequently furnished written sequence listing do stilled has been furnished.		
Certain claims were four Unity of invention is lack	nd unsearchable (See Box I). king (see Box II).		
4. With regard to the title,			
the text is approved as sul	omitted by the applicant.		
— ` ` ` `	ned by this Authority to read as follows:		
5. With regard to the abstract, X the text is approved as substract the text has been establish within one month from the 6. The figure of the drawings to be publicated by the application of the decause the applicant faile	ned, according to Rule 38.2(b), by this Authority date of mailing of this international search reposite with the abstract is Figure No.	y as it appears in Box III. The applicant may, ort, submit comments to this Authority. None of the figures.	
because this figure better of	characterizes the invention.		



International Application No PCT/US 01/29636

A. CLASSI IPC 7	FICATION OF SUBJECT MATTER A61K31/715 A61K39/395 A61P31/	04	
According to	o International Patent Classification (IPC) or to both national classific	cation and IPC	
B. FIELDS	SEARCHED		
Minimum do IPC 7	ocumentation searched (classification system followed by classification $A61K - A61P$	tion symbols)	
	tion searched other than minimum documentation to the extent that		
	ata base consulted during the international search (name of data ba	•))
BIOSIS	, PAJ, EMBASE, EPO-Internal, WPI Da	ta	
C. DOCUME	ENTS CONSIDERED TO BE RELEVANT		
Category °	Citation of document, with indication, where appropriate, of the re	elevant passages	Relevant to claim No.
X	SCHRAGER HARRY M ET AL: "Hyalur capsule modulates M protein-mediadherence and acts as a ligand fattachment of group A Streptococ CD44 on human keratinocytes." JOURNAL OF CLINICAL INVESTIGATION vol. 101, no. 8, 15 April 1998 (1998-04-15), page 1708-1716, XP002191017 ISSN: 0021-9738 abstract page 1711, column 1, paragraph 1 1713, column 2, paragraph 2 figure 3; table 2	ated or cus to N,	1-72
χ Furth	er documents are listed in the continuation of box C.	Patent family members are listed in	n annex.
° Special cat	egories of cited documents :	*T* later document published after the inter or priority date and not in conflict with	mational filing date
"E" earlier difiling da "L" documer which is citation "O" documer other m	ered to be of particular relevance ocument but published on or after the international ate at which may throw doubts on priority claim(s) or s citled to establish the publication date of another or other special reason (as specified) and referring to an oral disclosure, use, exhibition or seans	cited to understand the principle or the invention "X" document of particular relevance; the cl cannot be considered novel or cannot involve an inventive step when the document of particular relevance; the cl cannot be considered to involve an inv document is combined with one or moments, such combination being obviou	almed invention be considered to urment is taken alone almed invention entive step when the re other such docu-
later tha		in the art. *&* document member of the same patent f	amily
	ctual completion of the international search February 2002	Date of mailing of the international sear	rch report
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ivame and m	ailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Authorized officer Didelon, F	



International Application No PCT/US 01/29636

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.	
Х	PATENT ABSTRACTS OF JAPAN vol. 018, no. 385 (C-1227), 20 July 1994 (1994-07-20) & JP 06 107550 A (YUTAKA KURACHI), 19 April 1994 (1994-04-19) abstract	73-90	
(PATENT ABSTRACTS OF JAPAN vol. 018, no. 140 (C-1177), 8 March 1994 (1994-03-08) & JP 05 320055 A (DENKI KAGAKU KOGYO KK), 3 December 1993 (1993-12-03) abstract	73-90	
P, X	CYWES COLETTE ET AL: "CD44 as a receptor for colonization of the pharynx by group A Streptococcus." JOURNAL OF CLINICAL INVESTIGATION, vol. 106, no. 8, October 2000 (2000-10), pages 995-1002, XP002191018 ISSN: 0021-9738 abstract page 999, column 1, paragraph 3 -page 1000, column 1, paragraph 2 figures 4-6	1-72	
Т	CYWES COLETTE ET AL: "Group A Streptococcus tissue invasion by CD44-mediated cell signalling." NATURE (LONDON), vol. 414, no. 6864, 2001, pages 648-652, XP002191019 6 December, 2001 ISSN: 0028-0836 abstract	1-72	

NTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No PCT/US 01/29636

Patent document cited in search report		Publication date		Patent family member(s)	Publication date
JP 06107550	Α	19-04-1994	NONE		
JP 05320055	Α	03-12-1993	NONE		